1/11

# HPV 31 L1 nucleotide sequence alignment.

31 L1 wt	( 1)	ATGTCTCTGTGGCGGCCTAGCGAGGCTACTGTCTACTTACCACCTGTCCC
31 L1 partial	( 1)	
31 L1 total	( 1)	TA.AATCTAC
31 L1 wt	( 51)	AGTGTCTAAAGTTGTAAGCACGGATGAATATGTAACACGAACCAACATAT
31 L1 partial	(51)	
31 L1 total	( 51)	CGCCTCTCCCCCA
31 L1 wt	( 101)	ATTATCACGCAGGCAGTGCTAGGCTGCTTACAGTAGGCCATCCAT
	( 101)	AT TATE CACCACACACTAC MANAGEMENT COMMENT
31 L1 partial	( 101)	.CCTTTCATT.GCCTCC
31 L1 total	( 101)	,0.,0
31 L1 wt	( 151)	TCCATACCTAAATCTGACAATCCTAAAAAAATAGTTGTACCAAAGGTGTC
31 L1 partial	(151)	
31 L1 total	(151)	TCAG
		·
31 L1 wt	( 201)	AGGATTACAATATAGGGTATTTAGGGTTCGTTTACCAGATCCAAACAAA
31 L1 partial	(201)	
31 L1 total	(201)	TTGCACACA.AGCG.
31 L1 wt		TTGGATTTCCTGATACATCTTTTTATAATCCTGAAACTCAACGCTTAGTT
31 L1 partial	( 251)	
31 L1 total	( 251)	.CTCACC
31 L1 wt	(301)	TGGGCCTGTGTTGGTTTAGAGGTAGGTCGCGGGCAGCCATTAGGTGTAGG
31 L1 wc		
31 L1 total	( 301)	TCGACA.ATAGC
SI LI COCAT	( 501)	
31 L1 wt	( 351)	TATTAGTGGTCATCCATTATTAAATAAATTTGATGACACTGAAAACTCTA
31 L1 partial	(351)	
31 L1 total		
31 L1 wt	( 401)	
31 L1 partial	( 401)	
31 L1 total	( 401)	.CCTATCCCA

FIG.1A

31 L1 wt	( 451)	GATTATAAACAAACACAACTGTGTTTACTTGGTTGCAAACCACCTATTGG
31 L1 partial	( 451)	
31 L1 total	( 451)	CCGCTGT.GTGAC
31 L1 wt	( 501)	AGAGCATTGGGGTAAAGGTAGTCCTTGTAGTAACAATGCTATTACCCCTG
31 L1 partial	(501)	
31 L1 total	( 501)	TACGTCATCCCA.
31 L1 wt	( 551)	GTGATTGTCCTCCATTAGAATTAAAAAATTCAGTTATACAAGATGGGGAT
31 L1 partial	(551)	
31 L1 total	( 551)	CAGGGCTCCCTC
01.11.4	( (01)	**************************************
31 L1 wt		ATGGTTGATACAGGCTTTGGAGCTATGGATTTTACTGCTTTACAAGACAC
31 L1 partial		
31 L1 total	( 601)	CCTCT
31 L1 wt	( 651)	TAAAAGTAATGTTCCTTTGGACATTTGTAATTCTATTTGTAAATATCCAG
31 L1 partial	(651)	
31 L1 total	( 651)	CGTCCACCGC
31 L1 wt	( 701)	ATTATCTTAAAATGGTTGCTGAGCCATATGGCGATACATTATTTTTTAT
31 L1 partial	(701)	
31 L1 total	(701)	.CCT.GGCACCCGCC
31 L1 wt	•	TTACGTAGGGAACAAATGTTTGTAAGGCATTTTTTTAATAGATCAGGCAC
31 L1 partial		GAGC
31 L1 total	(751)	GAGC
31 LI COCAI	( /51)	d
31 L1 wt	•	
•		CATACC.GCG
31 L1 total	( 801)	CATACC.GCG
31 L1 wt	( 851)	CAGCTACTTTAGCTAACAGTACATACTTTCCTACACCTAGCGGCTCCATG
31 L1 partial	(851)	.CCC.GTCCCCATATCT
31 L1 total	( 851)	
31 L1 wt	( 901)	GTTACTTCAGATGCACAAATTTTTAATAAACCATATTGGATGCAACGTGC
-	•	CCCTGCCGC
31 L1 partial		
31 L1 total	(901)	CCCTGCCGC

31 L1 wt 31 L1 partial 31 L1 total	( 951) ( 951) ( 951)	TCAGGGACACAATAATGGTATTTGTTGGGGCAATCAGTTATTTGTTACTG ATCCTCCGCG ATCCTCCGCG
31 L1 wt 31 L1 partial 31 L1 total	(1001) (1001) (1001)	TGGTAGATACCACACGTAGTACCAATATGTCTGTTTGTGCTGCAATTGCA          C.        C.        C.        C.        C.        C.        C.        C.        T
31 L1 wt 31 L1 partial 31 L1 total	(1051) (1051) (1051)	AACAGTGATACTACATTTAAAAGTAGTAATTTTAAAGAGTATTTAAGACATCCCGTCCTCCGCC.GTCCCGTCCTCCGCC.G
31 L1 wt 31 L1 partial 31 L1 total	(1101) (1101) (1101)	TGGTGAGGAATTTGATTTACAATTTATATTTCAGTTATGCAAAATAACAT
31 L1 wt 31 L1 partial 31 L1 total	(1151) (1151) (1151)	TATCTGCAGACATAATGACATATTTCACAGTATGAATCCTGCTATTTTG .GTCCCCCCCCC
31 L1 wt 31 L1 partial 31 L1 total	(1201) (1201) (1201)	GAAGATTGGAATTTTGGATTGACCACACCTCCCTCAGGTTCTTTGGAGGAGCCTCTATC
31 L1 wt 31 L1 partial 31 L1 total	(1251) (1251) (1251)	TACCTATAGGTTTGTAACCTCACAGGCCATTACATGTCAAAAAAGTGCCC CC.A.C.C.T.A.T.C.C.GTCT.
31 L1 wt 31 L1 partial 31 L1 total		CCCAAAAGCCCAAGGAAGATCCATTTAAAGATTATGTATTTTGGGAGGTT .AACCGCCCAC
31 L1 wt 31 L1 partial 31 L1 total	(1351) (1351) (1351)	AATTTAAAAGAAAAGTTTTCTGCAGATTTAGATCAGTTTCCACTGGGTCGCGG
31 L1 wt 31 L1 partial 31 L1 total	(1401)	CAAATTTTATTACAGGCAGGATATAGGGCACGTCCTAAATTTAAAGCAG AGCGGATTCATA.AAGCGT.

PCT/US2004/008677

## 4/11

31 L1 partial	(1451)	GTAAACGTAGTGCACCCTCAGCATCTACCACTACACCAGCAAAACGTAAAGA.ATCTATTCCTGA.AG
31 L1 partial	(1501)	AAAACTAAAAAGTAA (SEQ ID NO:1) (SEQ ID NO:2) (SEQ ID NO:3)

FIG.1D

HPV31 L1 total rebuild nucleotide and amino acid sequences.

- M S L W R P S E A T V Y L P P V P

  1 ATGTCTTTGT GGAGACCATC TGAAGCTACC GTCTACTTGC CACCAGTCCC
- V S K V V S T D E Y V T R T N I Y 51 AGTCTCTAAG GTCGTCTCTA CCGACGAATA CGTCACCAGA ACCAACATCT
- Y H A G S A R L L T V G H P Y Y

  101 ACTACCACGC TGGTTCTGCT AGATTGTTGA CCGTCGGTCA CCCATACTAC
- S I P K S D N P K K I V V P K V S

  151 TCTATCCCAA AGTCTGACAA CCCAAAGAAG ATCGTCGTCC CAAAGGTCTC
- G L Q Y R V F R V R L P D P N K F 201 TGGTTTGCAA TACAGAGTCT TCAGAGTCAG ATTGCCAGAC CCAAACAAGT
- G F P D T S F Y N P E T Q R L V
  251 TCGGTTTCCC AGACACCTCT TTCTACAACC CAGAAACCCA AAGATTGGTC
- W A C V G L E V G R G Q P L G V G 301 TGGGCTTGTG TCGGTTTGGA AGTCGGTAGA GGTCAACCAT TGGGTGTCGG
- I S G H P L L N K F D D T E N S N 351 TATCTCTGGT CACCCATTGT TGAACAAGTT CGACGACACC GAAAACTCTA
- R Y A G G P G T D N R E C I S M 401 ACAGATACGC TGGTGGTCCA GGTACCGACA ACAGAGAATG TATCTCTATG
- D Y K Q T Q L C L L G C K P P I G 451 GACTACAAGC AAACCCAATT GTGTTTGTTG GGTTGTAAGC CACCAATCGG
- E H W G K G S P C S N N A I T P G 501 TGAACACTGG GGTAAGGGTT CTCCATGTTC TAACAACGCT ATCACCCCAG
- D C P P L E L K N S V I Q D G D 551 GTGACTGTCC ACCATTGGAA TTGAAGAACT CTGTCATCCA AGACGGTGAC

# FIG. 2A

601	M V D T G F G A M D F T A L Q D T ATGGTCGACA CCGGTTTCGG TGCTATGGAC TTCACCGCTT TGCAAGACAC
651	K S N V P L D I C N S I C K Y P D CAAGTCTAAC GTCCCATTGG ACATCTGTAA CTCTATCTGT AAGTACCCAG
701	Y L K M V A E P Y G D T L F F Y ACTACTTGAA GATGGTCGCT GAACCATACG GCGACACCTT GTTCTTCTAC
751	LRRE QMF VRH FFNRS GT TTGCGTAGAG AACAGATGTT CGTAAGGCAC TTCTTCAACA GATCCGGCAC
801	V G E S V P T D L Y I K G S G S T CGTAGGTGAA TCTGTCCCAA CCGACCTGTA CATCAAGGGC TCCGGTTCCA
851	A T L A N S T Y F P T P S G S M CCGCTACCCT GGCTAACTCC ACCTACTTCC CAACTCCATC TGGCTCCATG
901	V T S D A Q I F N K P Y W M Q R A GTCACCTCCG ACGCTCAGAT CTTCAACAAG CCATACTGGA TGCAGCGTGC
951	Q G H N N G I C W G N Q L F V T V ACAGGGTCAC AACAACGGTA TCTGTTGGGG TAACCAGCTG TTCGTGACTG
1001	V D T T R S T N M S V C A A I A TGGTCGATAC CACGCGTTCT ACCAACATGT CTGTCTGTGC TGCAATCGCT
1051	N S D T T F K S S N F K E Y L R H AACTCTGACA CTACCTTCAA GTCCTCTAAC TTCAAGGAGT ACCTGAGACA
1101	G E E F D L Q F I F Q L C K I T L TGGTGAGGAA TTCGATCTGC AATTCATCTT CCAGTTGTGC AAGATCACCC
1151	S A D I M T Y I H S M N P A I L TGTCTGCTGA CATCATGACC TACATCCACA GTATGAACCC TGCCATCCTG
1201	E D W N F G L T T P P S G S L E D GAGGACTGGA ACTTCGGTCT GACCACTCCA CCTTCCGGTT CTTTGGAAGA

# FIG.2B

	T	Υ	R	F	٧	T	S	Q	F	1	I	T	C	(	2	K	S	Α	P
1251	CAC	CTA	CAGA	TTO	CGT	CACC	T	CTC	AA(	GCT	TAT	CAC	CCT	GT(	CAA	AA	GTC'	TGC	TC
			Р																
1301	CAC	AAA	AGCC	AAA	AGG/	AAGA	4C	CCA	ПС	ÇΑΡ	AGG	AC	IAC	GI(	311	CI	GGG.	AAG	IC
	N i	ı	K E	L	, ;	E (	2	Λ	ח	1		n	0	E	D	!	1 1	<b>G</b> 1	D
1351																			
1001	7010	, . u	, u iaa	,,,,	<i>-</i> 10					ιο.		<b></b>				0, .		<b></b>	
	Κ	F	L	L	Q	Α	G	Υ	F	₹	Α	R	Р	ŀ	<	F	K	Α	G
1401	AAA	GTT	CTTG	TTO	GCA	AGCT	ΓG	GTT	AC/	٩GA	AGC	TAG	<b>GAC</b>	CA/	4AG	TT	CAA	GGC'	TG
												_	_						
			S																
1451	GTA	AGA	GATC	TG(	CTC	CATO	CT	GCT	TCT	ΓAC	CA	CCA	ACC	CCA	AGC	TA	AGA	GAA	AG
	<i>v</i> -	т	K K	,	k (1	CEU	TF	NO	. 4										
1 - 0 1																			
1501	AAG	AUU	AAGA	Au	IAA	(SE	:Ų	ıυ	NU :	:3)	,								

FIG.2C

SUMMARY OF HPV 31 SEQUENCES.

			z
COMMENTS		121 CHANGES BETWEEN nt 697-1249	376 CHANGES BETWEEN nt 1-1515
AMINO ACID IDENTITY		100%	100%
ACIDS NUCLEOTIDE IDENTITY AMINO ACID IDENTITY		92%	75%
1 1	504	504	504
NUCLEOTIDES	1515	1515	1515
L1 CONSTRUCT NUCLEOTIDES AMINO	HPV 31 LA WILD-TYPE	HPV 31 L1 PARTIAL REBUILD	HPV 31 L1 TOTAL REBUILD

FIG.3

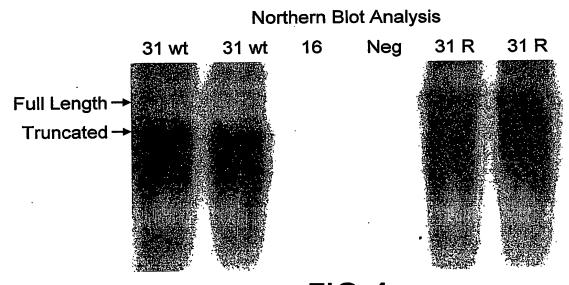


FIG.4

RESULTS FROM RADIOIMMUNOASSAY

10/11

RATIO (Ave/Ave)	(31 wt/31 wt)	6.9 (31 PARTIAL/31 wt)	1 (31 Partial/31 Partial)	1 (31 TOTAL/31 PARTIAL)
AVERAGE (cpm/mg)	460	3158	2095	3645
31 L1 VLP*/ mg protein (cpm/mg)	252 609 497 480	3031 3285	1719 1798 2998 1861	3202 4087
RIA MINUS BACKGRUOND 31 L1 VLP*/ mg protein (cpm/ml) (cpm/mg)	414 987 904 844	5061 5091	3901 4081 7135 4428	6628 8461
PROTEIN CONCENTRATION (mg/ml)	1.64 1.62 1.82 1.76	) 1.67. 1.55	D 2.27 2.27 2.38 2.38	2.07
EXP. L1 CONSTRUCT CO	31 WILD-TYPE	31 Partial Rebuild	31 Partial Rebuild	31 TOTAL REBUILD
EXP.	-	<del></del>	8	2

FIG. 5

WO 2004/084831 PCT/US2004/008677

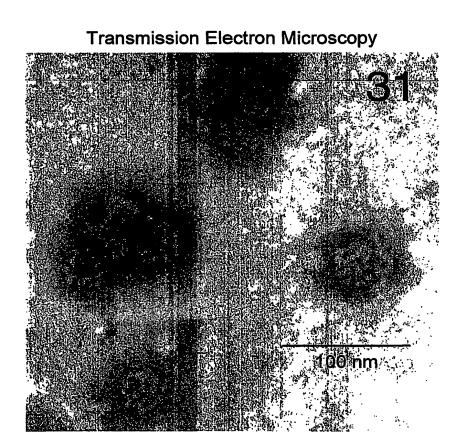


FIG.6